

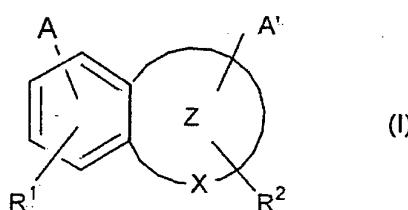
APPLICANT(S): ZIV, Ilan et al.
SERIAL NO.: Not yet assigned
FILED: Herewith
Page 9

AMENDMENTS TO THE CLAIMS

Please cancel claims 1-41. Please add new claims 42-67 as follows.

Claims 1-41 (Cancelled)

42. (New) A method for selective targeting of a chemical compound to a cell undergoing perturbation of the *normal organization* of its plasma membrane (PNOM-cell) present in a cell population, comprising the steps of: (i) contacting the cell population with a PMBC, being a chemical compound represented by the structure set forth in formula (I):



wherein Z represents null, or a ring system formed of cycloalkyl, cycloalkenyl, heterocyclyl, aryl or heteroaryl groups or combinations of such groups, the ring system consisting of 5, 6, 7, 8, 9 or 10 atoms;

X represents an atom, which is C, N, O or S, where each of these atoms may bear 0, 1 or 2 hydrogen atoms according to the meaning of Z;

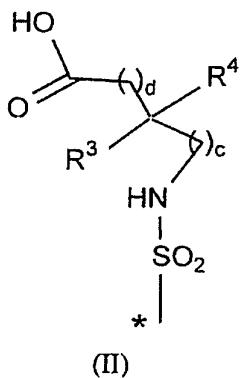
R¹ and R² are each independently hydrogen, halogen, hydroxyl, -NO₂ group or W-Q_b; wherein W is null, nitrogen, oxygen or carbon; and Q represents hydrogen, a C₁, C₂, C₃, C₄, C₅ or C₆ alkyl, hydroxyalkyl, or straight or branched haloalkyl, wherein Q groups may be either the same or different; and b is an integer, being 1 when W is oxygen or null; 2 in the case that W is nitrogen; or 3 in the case that W is a carbon atom;

A and A' are each a radical independently selected from one of the following four groups:

- i) hydrogen;
- ii) SO₃H, and L-SO₃H, wherein L stands for a C₁, C₂, C₃, C₄ or C₅ alkylene linker;

APPLICANT(S): ZIV, Ilan et al.
SERIAL NO.: Not yet assigned
FILED: Herewith
Page 10

iii) a structure, set forth in formula II:

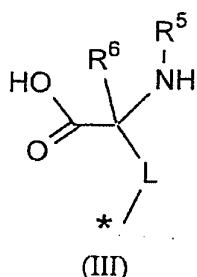


wherein R³ is hydrogen, (CH₂)_p-OH, (CH₂)_p-SH, (CH₂)_p-F, or suspected of comprising such PNOM cell, comprising the steps of contacting a cell population with a compound represented by a radical of C₁, C₂, C₃, or C₄ carboxylic acid, wherein p is 1, 2, or 3;

R⁴ is hydrogen, a C₁, C₂, C₃, C₄, C₅ C₆ straight or branched alkyl, a C₁, C₂, C₃, C₄, C₅ C₆ straight or branched hydroxyalkyl or a C₁, C₂, C₃, C₄, C₅ C₆ straight or branched fluoroalkyl;

c and d are each an integer of 0 or 1; c and d may be the same or different; * represents the point of attachment to the structure of formula (I); or

iv) a structure set forth in formula (III):



wherein R⁵ and R⁶ are independently hydrogen, C₁, C₂, C₃, C₄, C₅, C₆ straight or branched alkyl, C₁, C₂, C₃, C₄, C₅, C₆ straight or branched hydroxyalkyl or C₁, C₂, C₃, C₄, C₅, C₆ straight or branched haloalkyl; R⁵ and R⁶ can be the same or different; and L stands for null

APPLICANT(S): ZIV, Ilan et al.
SERIAL NO.: Not yet assigned
FILED: Herewith
Page 11

or a C₁, C₂, C₃, C₄ or C₅ alkylene linker; * represents the point of attachment to the structure of formula (I);

wherein when at least one of A or A' groups is other than hydrogen, and is different than the structure of formula (III);

thereby selectively targeting the chemical compound to the PNOM-cells within the cell population.

43. (New) A method of detecting the presence of PNOM-cells within a cell population, comprising the steps of:

(i) administrating the cell population with a PMBC, or a conjugate comprising said PMBC and a marker for imaging, wherein said PMBC is represented by the structure set forth in formula (I) of Claim 42, wherein A, A', X, Z, R¹, R², R³, R⁴, R⁵, R⁶, L, c and d are as defined in Claim 42; and

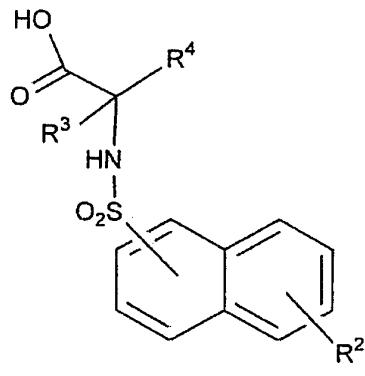
(ii) determining the amount of PMBC bound to cells in the cell population wherein a bound amount which is significantly higher than a control indicates the presence of PNOM-cells within the cell population.

44. (New) A method according to Claim 42, wherein the PNOM- cell is a cell undergoing a death process, an apoptotic cell or an activated platelet.

45. (New) A method according to Claim 42, wherein in the compound represented by the structure as set forth in formula (I) A or A' are represented by formula (II), R¹ is hydrogen, and R² is NQ₂, wherein Q groups may be the same or different, each being a hydrogen or a C₁-C₄ alkyl.

46. (New) A method according to Claim 42, wherein said PMBC is represented by the structure as set forth in formula (IV):

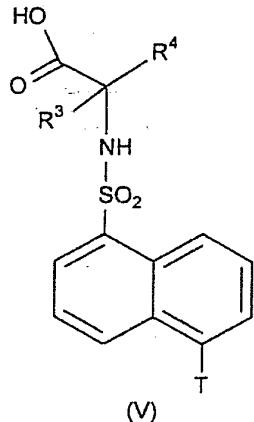
APPLICANT(S): ZIV, Ilan et al.
SERIAL NO.: Not yet assigned
FILED: Herewith
Page 12



(IV)

wherein R², R³ and R⁴ are as defined in Claim 42.

47. (New) A compound represented by a structure as set forth in formula (V):

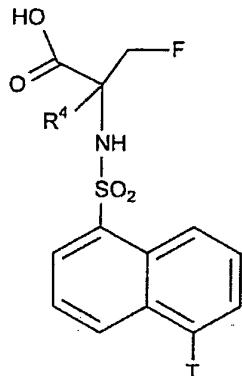


(V)

wherein T is -OH, -O-CH₃, -O-(CH₂)_yCH₃, NH₂, N(CH₃)₂, N[(CH₂)₃CH₃]₂, -N(CH₃)[(CH₂)₂CH₃], -N(CH₃)CH₂CH₃ or -N(CH₃)[(CH₂)₃CH₃]; y stands for an integer of 1, 2, or 3; and R³ and R⁴ are each as defined in Claim 42.

APPLICANT(S): ZIV, Ilan et al.
SERIAL NO.: Not yet assigned
FILED: Herewith
Page 13

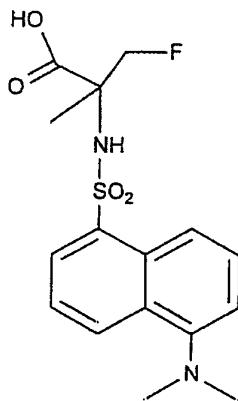
48.(New) A compound according to Claim 47, represented by the structure as set forth in formula (VI):



(VI)

wherein T is as defined in Claim 47, and R⁴ is hydrogen or a C₁, C₂, C₃, C₄, C₅ or C₆ straight or branched alkyl, and wherein the F atom is ¹⁸F or ¹⁹F or mixtures of fluorine isotopes.

49. (New) A compound according to Claim 47, represented by the structure as set forth in formula (VII):

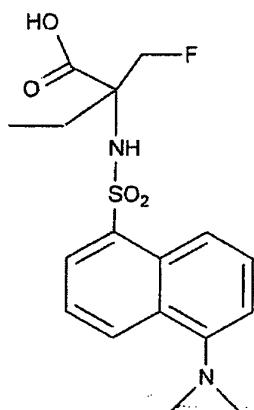


(VII)

APPLICANT(S): ZIV, Ilan et al.
SERIAL NO.: Not yet assigned
FILED: Herewith
Page 14

wherein the F atom is ^{18}F or ^{19}F or mixture of fluorine isotopes.

50. (New) A compound according to Claim 47, represented by the structure as set forth in formula (VIII):

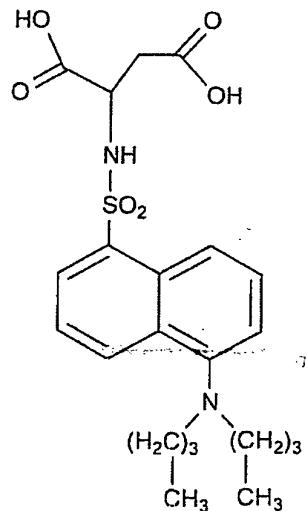


(VIII)

wherein the F atom is ^{18}F or ^{19}F or mixtures of fluorine isotopes.

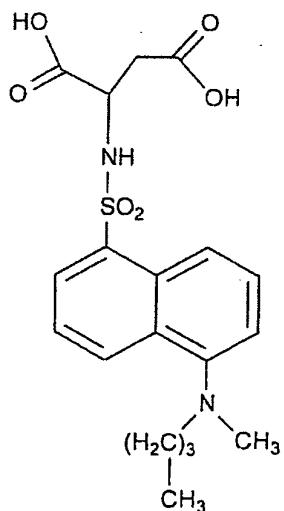
51. (New) A compound according to Claim 47 represented by the structure as set forth in formula (IX):

APPLICANT(S): ZIV, Ilan et al.
SERIAL NO.: Not yet assigned
FILED: Herewith
Page 15



(IX)

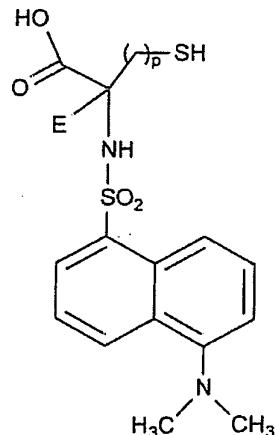
52. (New) A compound according to Claim 47, represented by the structure as set forth in formula (X):



(X)

APPLICANT(S): ZIV, Ilan et al.
SERIAL NO.: Not yet assigned
FILED: Herewith
Page 16

53.(New) A compound according to Claim 47, represented by the structure as set forth in formula (XI):

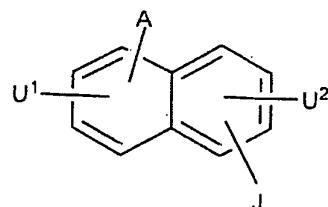


(XI)

wherein E is C₁, C₂, C₃ or C₄ alkyl; C₁, C₂, C₃ or C₄ fluoroalkyl; or C₁, C₂, C₃ or C₄ hydroxyalkyl; p stands for an integer of 1 or 2.

54. (New) A compound according to Claim 53, wherein p is 1.

55. (New) A method according to claim 42 comprising the step of contacting a cell population with a compound represented by the structure as set forth in formula (XII):



(XII)

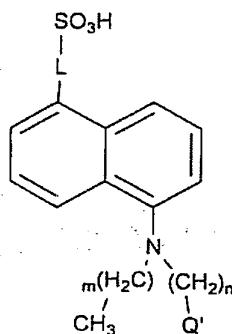
APPLICANT(S): ZIV, Ilan et al.
SERIAL NO.: Not yet assigned
FILED: Herewith
Page 17

wherein A is SO_3H or L- SO_3H , wherein L stands for a substituted or unsubstituted C_1 , C_2 , C_3 , C_4 or C_5 alkylene;

J is selected from SO_3H , L- SO_3H , wherein L is as defined above, hydrogen and W-Q_b; wherein W is null, nitrogen, oxygen or carbon; and Q represents hydrogen, a C_1 , C_2 , C_3 , C_4 , C_5 or C_6 straight or branched alkyl, straight or branched hydroxyalkyl, or straight or branched haloalkyl; wherein Q groups may be either the same or different; b is an integer, being 1 when W is oxygen or null, 2 in the case that W is nitrogen, or 3 in the case that W is a carbon atom;

U^1 and U^2 are each halogen, hydrogen, hydroxyl, $-\text{NO}_2$; C_1 , C_2 , C_3 , C_4 , C_5 or C_6 straight or branched alkyl; C_1 , C_2 , C_3 , C_4 , C_5 or C_6 straight or branched halo-alkyl; C_1 , C_2 , C_3 , C_4 , C_5 or C_6 straight or branched hydroxy-alkyl; U^1 and U^2 groups may be the same or different.

56. (New) A method according to claim 55, comprising the step of contacting a cell population with a compound represented by the structure as set forth in formula (XIII):

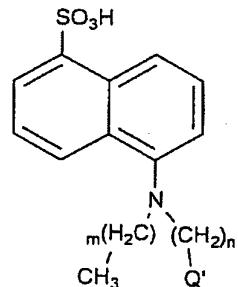


(XIII)

wherein n stands for an integer of 1, 2, 3, 4, 5 or 6, m stands for an integer of 0, 1, 2 or 3, Q' is hydrogen, $-\text{OH}$ or $-\text{F}$, and L stands for null or C_1 , C_2 , C_3 , C_4 or C_5 alkylene, thereby selectively targeting a PNOM-cell in said cell population.

57. (New) A method according to claim 55, wherein said cell population is suspected of comprising such a PNOM-cell, comprising the step of contacting a cell population with a compound represented by the structure as set forth in formula (XIV);

APPLICANT(S): ZIV, Ilan et al.
SERIAL NO.: Not yet assigned
FILED: Herewith
Page 18



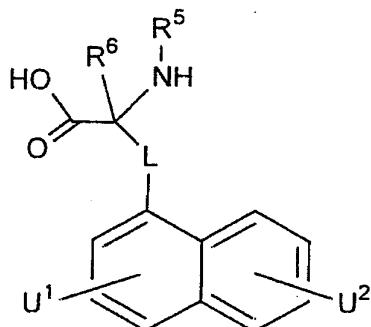
(XIV)

wherein n stands for an integer of 1, 2, 3, 4, 5 or 6, m stands for an integer of 0, 1, 2 or 3 and Q' is hydrogen, $-\text{OH}$ or $-\text{F}$.

58. (New) A compound represented by the structure set forth in formula (XIV) wherein Q' is F, either ^{18}F or ^{19}F or a mixture of isotopes or wherein m is 0, n is 4 and Q' is hydrogen or wherein m is 0, n is 3 and Q' is hydroxyl or wherein m is 0, n is 4 and Q' is fluorine.

59. (New) A method according to claim 57, comprising the step of contacting a cell population with a compound represented by the structure as set forth in formula (XV):

APPLICANT(S): ZIV, Ilan et al.
SERIAL NO.: Not yet assigned
FILED: Herewith
Page 19



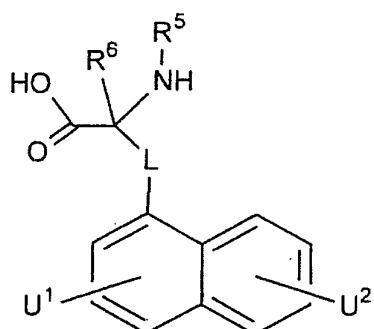
(XV)

wherein R⁵ and R⁶ are each independently hydrogen, C₁, C₂, C₃, C₄, C₅, C₆ straight or branched alkyl, straight or branched hydroxyalkyl or straight or branched fluoroalkyl; R⁵ and R⁶ can be the same or different;

L stands for null or a C₁, C₂, C₃, C₄ or C₅ alkylene linker;

U¹ and U² are each hydrogen, halogen, hydroxyl, -NO₂; C₁, C₂, C₃, C₄, C₅ or C₆ straight or branched alkyl; C₁, C₂, C₃, C₄, C₅ or C₆ straight or branched haloalkyl; C₁, C₂, C₃, C₄, C₅ or C₆ straight or branched hydroxyalkyl; U groups may be the same or different.

60. (New) A compound represented by the structure as set forth in formula (XV),

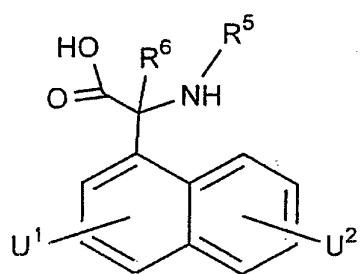


(XV)

APPLICANT(S): ZIV, Ilan et al.
SERIAL NO.: Not yet assigned
FILED: Herewith
Page 20

wherein either U^1 or U^2 is fluorine or C_1, C_2, C_3 or C_4 fluoroalkyl, the F atom being either ^{18}F or ^{19}F .

61. (New) A compound represented by the structure as set forth in formula (XVI):



(XVI)

wherein R^5 and R^6 are independently hydrogen, $C_1, C_2, C_3, C_4, C_5, C_6$ straight or branched alkyl, straight or branched hydroxyalkyl or straight or branched fluoroalkyl; R^5 and R^6 can be the same or different; U^1 and U^2 are each hydrogen, halogen, hydroxyl, $-NO_2$; C_1, C_2, C_3, C_4, C_5 or C_6 straight or branched alkyl; C_1, C_2, C_3, C_4, C_5 or C_6 straight or branched haloalkyl; C_1, C_2, C_3, C_4, C_5 or C_6 straight or branched hydroxyalkyl; U groups may be the same or different.

62. (New) A compound according to claim 61 represented by the structure as set forth in formula (XVI), wherein R^5 is $C_1, C_2, C_3, C_4, C_5, C_6$ straight or branched alkyl, straight or branched hydroxyalkyl and straight or branched fluoroalkyl; R^6 is C_2, C_3, C_4, C_5, C_6 straight or branched alkyl, straight or branched hydroxyalkyl or straight or branched fluoroalkyl; and either U^1 or U^2 is other than hydrogen.

APPLICANT(S): ZIV, Ilan et al.
SERIAL NO.: Not yet assigned
FILED: Herewith
Page 21

63. (New) A method of selective targeting PNOM-cells comprising the step of contacting a cell population with a compound represented by the structure as set forth claim 61.

64. (New) A method according to claim 43, comprising the steps of:

(i) contacting the cell population with a PMBC, or a conjugate comprising said PMBC and a marker for imaging, wherein said PMBC is represented by the structure set forth represented in any one of formulae I, IV, V, VI, VII, VIII, IX, X, XI, XII, XIII, XIV, XV or XVI; and

(II) determining the amount of PMBC bound to cells in said cell population, wherein a bound amount which is significantly higher than a control indicates the presence of PNOM-cells within the cell population.

65. (New) A compound according to Claim 47 wherein said compound is being linked either directly or through a linker Y to a member selected from a solid support, a marker for imaging or a therapeutic drug, wherein said linker Y is C₁, C₂, C₃, C₄, C₅ or C₆ alkylene, 5-6 atom aromatic or 5-6 heteroaromatic ring, wherein the heteroatom of said heteroaromatic ring is N, O and S, a metal chelator, or combinations thereof.

66. (New) An agent for the detection of PNOM-cells, comprising a compound according to claim 47, wherein said compound is linked or comprises a marker for imaging.

67. (New) An agent according to claim 66 wherein said marker for imaging is detectable by detector of color, fluorescence, X-ray, CT scan, MRI, radio-isotope scan, SPECT, or PET scan.